

**BIOGRAPHICAL SKETCH**

Provide the following information for the key personnel in the order listed for Form Page 2.  
Follow the sample format on preceding page for each person. DO NOT EXCEED FOUR PAGES.

NAME		POSITION TITLE	
Adam Lerner		Associate Professor of Medicine	
EDUCATION/TRAINING <i>(Begin with baccalaureate or other initial professional education, such as nursing, and include postdoctoral training.)</i>			
INSTITUTION AND LOCATION	DEGREE <i>(if applicable)</i>	YEAR(s)	FIELD OF STUDY
Amherst College	A.B.	1979	Biology
Yale Medical School	M.D.	1983	Medicine

**A. Positions and Honors:**Internship and Residencies

1983-1984 Intern in Internal Medicine, Boston City Hospital, Boston, MA

1984-1986 Resident in Internal Medicine, Boston City Hospital

Research and Clinical Positions

1986-1988 Research Fellow, Department of Pathology, Boston University School of Medicine, Boston, MA

1988-1991 Clinical Fellow in Oncology, Dana-Farber Cancer Institute, Harvard Medical School, Boston, MA

1991-1993 Research Fellow, Laboratory of Immunobiology, Dana-Farber Cancer Institute, Boston, MA

1993-1995 Instructor, Department of Medicine, Harvard Medical School, Boston, MA

1996- Assistant Professor, Depts. Of Medicine and Pathology, Boston University School of Medicine, Boston, MA

2002- Associate Professor, Depts. Of Medicine and Pathology, Boston University School of Medicine, Boston, MA

2000- Member, Immunology Training Program, Boston University School of Medicine, Boston, MA

2000- Member and Instructor, Graduate Program in Molecular Medicine, Boston Medical Center

2001- Chairman, Human Gene Therapy Committee, Boston University School of Medicine

2006- Principal Investigator, NHLBI Hematology Training Grant, BU School of Medicine, Boston, MA

Awards, Honors and Support:

1979 Phi Beta Kappa, magna cum laude (Amherst College)

1992-1995 Leukemia Society of America Special Fellowship

1997-1999 Leukemia Society of America Translational Research Award

1997-1998 American Cancer Society, Massachusetts Division, Research Award

1998-2001 American Society of Clinical Oncology Career Development Award

2000-2004 NCI RO1 Award: CA79838

2001-2004 DOD Breast Cancer Idea Award

2005-2010 NCI RO1 Awards: CA106705 and CA114094

Membership in Professional Societies and Professional Activities:

1995- American Society of Clinical Oncology (ABIM Recertified in Oncology 2004), 1997- American Society of

Hematology, 2002- American Association of Immunologists, American Association for Cancer Research.

2005- *ad hoc* NIH reviewer: ZRG1 IMM-B (Cell and Molecular Immunology)

2005, 2006- *ad hoc* NIH reviewer: ZRG1-ONC-Q (Basic Mechanisms of Cancer Therapeutics)

**B. Selected publications (in chronological order):**

1. Lerner A, Jacobson B, Miller R. Cyclic AMP concentrations modulate both calcium flux and hydrolysis of phosphatidylinositol phosphates in mouse T lymphocytes. *J Immunol*. 1988; 140:936-940.
2. Lerner A, Phillosophe B, Miller R. Defective calcium influx and preserved inositol phosphate generation in T cells in old mice. *Aging: Immunol & Infectious Disease* 1988; 1:149-157.
3. Lerner A, Yamada A, Miller R. PGP-1 T lymphocytes accumulate with age in mice and respond poorly to concanavalin A. *Eur J Immunol*. 1989; 19:977-982.
4. Lerner A, Gonin R, Steele GD, Mayer RJ. Etoposide, doxorubicin, and cisplatin chemotherapy for advanced gastric adenocarcinoma: results of a phase II trial. *J. Clin. Oncol*. 1992; 10:536-540.

5. **Lerner A**, Diener AC, Reinherz EL, Clayton LK. Human genomic sequences corresponding to murine CD3 $\eta$ -related transcripts: lack of expression of homologous human products. *Eur J Immunol.* 1992; 22:2135-2140.
6. **Lerner A**, D'Adamio L, Diener AC, Clayton LK and Reinherz EL. The CD3 $\zeta$ / $\eta$ / $\theta$  locus is colinear with and transcribed antisense to the gene encoding the transcription factor Oct-1. *J Immunol.* 1993; 151:3152-3162.
7. Clayton LK, Diener AC, **Lerner A**, Tse A, Koyasu S and Reinherz EL. Differential regulation of T-cell receptor processing and surface expression affected by CD3 $\eta$ , an alternatively spliced product of the CD3 $\zeta$ / $\eta$ / $\theta$  gene locus. *J Biol. Chem.* 1992; 267:26023-26030.
8. Clayton LK, **Lerner A**, Diener AC, Hussey RE, Koyasu S, and Reinherz EL. T cell receptor isoforms. *Int. J. Cancer* 1992; Supplement 7:1-5.
9. Reinherz EL, **Lerner A**, Diener AC, Hussey R, Koyasu S and Clayton LK. T cell receptor isoforms during thymic differentiation. *Mol. Basis of Immune Resp.*, H. Nariuchi ed. Academic Press, 1993.
10. Koyasu S, Hussey RE, Clayton LK, **Lerner A**, Pederson R, Delaney-Heiken P, Chau F and Reinherz EL. Targeted disruption within the CD3 $\zeta$ / $\eta$ / $\theta$  locus in mouse. *EMBO J.* 1994; 13:784-797.
11. **Lerner A**, Clayton LC, Mizoguchi E, Ghendler Y, Koyasu S, Bhan AK and Reinherz EL. Cross-linking of T cell receptors on double positive thymocytes induces a cytokine-mediated stromal activation process linked to cell death. *EMBO J.* 1996; 15:5876-5887.
12. Koyasu S, Clayton LC, **Lerner A**, Heiken H, Parkes A, Reinherz EL. Pre-TCR signaling components trigger transcriptional activation of a rearranged TCR $\alpha$  gene locus and silencing of the pre-TCR $\alpha$  locus: implications for intrathymic differentiation. *Int. Immunol.* 1997; 9:1475-1480.
13. Kim, D.H. and **Lerner A**. Type 4 cAMP phosphodiesterase as a therapeutic target in chronic lymphocytic leukemia. *Blood* 1998; Vol 92, No.7: 2484-2494.
14. Cai D, Clayton LK, Smolyar A and **Lerner A**. AND-34, a novel p130<sup>Cas</sup>-binding thymic stromal cell protein induced by inflammatory cytokines. *J. Immunol.* 1999; 163: 2104-2112.
15. **Lerner A**, Elias A. Bone and soft tissue sarcomas. *Current Practice in Medicine*, Vol. XII: 9.1-9.7. Churchill Livingstone Inc., New York, N.Y. 1996 (1<sup>st</sup> edition) and 1999 (2<sup>nd</sup> edition).
16. Gotoh T, Cai D, Feig L and **Lerner A**. P130<sup>Cas</sup> regulates the activity of AND-34, a novel Ral, Rap1 and R-Ras guanine nucleotide exchange factor. *J. Biol. Chem.* 2000; 275: 30118-30123.
17. Lee R, Kim DH and **Lerner A**. The cAMP signaling pathway as a therapeutic target in lymphoid malignancies. *Leukemia and Lymphoma*. 2000, Vol 37 (1-2): 39-51.
18. Akpek G, Lenz G, Lee SM, Sancharawala V, Wright DG, Colarusso T, Waraska K, **Lerner A**, Vosburgh E, Skinner M and Comenzo RL. Immunologic recovery after autologous blood stem cell transplantation in patients with AL-amyloidosis. *Bone Marrow Transplantation* 2001; 28: 1105-1109.
19. Lee R, Wolda S, Moon E, Esselstyn J, Hertel C and **Lerner A**. PDE7A is expressed in human B lymphocytes and is up-regulated by elevation of intracellular cAMP. *Cellular Signalling*, 2002: 227-287.
20. Moon E, Lee R, Near R, Weintraub L, Wolda S, and **Lerner A**. Inhibition of PDE3B variably augments PDE4 inhibitor-induced apoptosis in chronic lymphocytic leukemia. *Clin. Cancer Research* 2002, 8:589-595.
21. Moon E and **Lerner A**. Benzylamide sulindac analogues induce changes in cell shape, loss of microtubules and G2/M arrest in a CLL cell line and apoptosis in primary CLL cells. *Cancer Research* 2002, 62:5711-5719.
22. Cai D, Felekis K, Near R, Iyer A, O'Neill G, Seventer J, Golemis E, **Lerner A**. The GDP exchange factor AND-34 is expressed in B cells, associates with HEF1, and activates Cdc42. *J. Immunol.* 2003, 170:969-978.
23. Cai D, Iyer A, Near R, Felekis KN, Luo Z, Albanese C, Pestell RG and **Lerner A**. AND-34/BCAR3, a GDP exchange factor whose over-expression confers antiestrogen resistance, activates Rac1, Pak1 and the cyclin D1 promoter. *Cancer Research* 2003, 63:6802-6808.
24. Moon E and **Lerner A**. PDE4 inhibitors activate a mitochondrial apoptotic pathway in chronic lymphocytic leukemia that is regulated by PP2A. *Blood* 2003; 101:4122-4130.
25. Tiwari S, Felekis K, Moon E-Y, Flies A, Sherr D, **Lerner A**. Among circulating hematopoietic cells, B-CLL uniquely expresses functional EPAC1 but EPAC1-Mediated Rap1 activation does not account for PDE4 inhibitor-induced apoptosis. *Blood* 2004; 103:2661-2667.
26. Koido S, Ohana M, Liu C, Nikrui N, Durfee J, **Lerner A** and Gong J. Dendritic cells fused with human cancer cells: Morphology, antigen expression and T cell stimulation. *Clin. Immunol.* 2004; 113:261-269.
27. **Lerner A** and Andrea N. Vinca alkaloids vs. taxanes as therapy in lymphoid malignancies: Do our experimental models obscure our knowledge of how these drugs really work? *Cancer Invest.* 2005;23:100-102.
28. Tiwari S, Dong H, Kim EJ, Weintraub L, Epstein PM and **Lerner A**. PDE4 inhibitors augment glucocorticoid-mediated apoptosis and signaling in B-CLL in the absence of exogenous adenylate cyclase stimulation. *Biochem. Pharmacol.* 2005; 69; 473-483.

29. Felekis KN, Narsimhan RP, Castro AF, Quilliam LA and **Lerner A**. AND-34 activates phosphatidylinositol 3-kinase and induces antiestrogen resistance in a SH2 and GEF-like domain-dependent manner. *Molecular Cancer Research*. 2005; 3; 32-41.
30. Koido S, Nikrui N, Ohana M, Xia J, Tanaka Y, Liu C, Durfee J, **Lerner A** and Gong J. Assessment of fusion cells from patient-derived ovarian carcinoma cells and dendritic cells as a vaccine for clinical use. *Gynecol. Oncol.* 2005; 99; 462-471.
31. **Lerner A** and Epstein P. Cyclic nucleotide phosphodiesterases as targets for treatment of haematological malignancies. *Biochemical Journal*. 2006; 393; 21-41.
32. **Lerner A** and Felekis KN. Analysis of AND-34-induced Rac and Cdc42 activation in lymphoid and epithelial cells. *Methods in Enzymology*, 2006; 407: 55-63.
33. **Lerner A**, Moon E and Tiwari S. Role of Phosphodiesterases in apoptosis. *Cyclic Nucleotide Phosphodiesterases in Health and Disease*, J Beavo, S Francis and M Houslay, ed. CRC Press, 2006. Pages 559-582.
34. Everett P, Meyers JA, Makkinje A, Rabbi M and **Lerner A**. Curcumin augments vinca alkaloid and PDE4 inhibitor-induced apoptosis in B-CLL cells at clinically tolerable concentrations. *Amer. J. Hematol.* 2007; 82: 23-30.
35. Taverna JA, **Lerner A**, Goldberg L, Werth S, Demierre MF. Infliximab as therapy for idiopathic hypereosinophilic syndrome. *Arch. Dermatol.* 2007; 143: 1110-1112.
36. Near RI, Zhang Y, Makkinje A, Vanden Borre P, and **Lerner A**. AND-34/BCAR3 differs from other NSP homologs in induction of anti-estrogen resistance, cyclin D1 promoter activation and altered breast cancer cell morphology. *J. Cell. Physiol.* 2007; 212: 655-665.
37. Taverna JA, **Lerner A**, Bhawan J, Demierre MF. Successful adjuvant treatment of recalcitrant mucous membrane pemphigoid with anti-CD20 antibody Rituximab. *J. Drugs Dermatol.* 2007; 6: 731-2.
38. Meyers JA, Taverna J, Chaves J, Makkinje A and **Lerner A**. PDE4 inhibitors augment levels of glucocorticoid receptor in B cell chronic lymphocytic leukemia but not in normal circulating hematopoietic cells. *Clinical Cancer Research* 2007; 13: 4920-4927.
39. Mineva ND, Rothstein TL, Meyers J, **Lerner A**, and Sonenshein GE. CD40 ligand mediated activation of the de novo RelB NF- $\kappa$ B synthesis pathway in transformed B cells promotes rescue from apoptosis. *J. Biol. Chem.* 2007; 282: 17475-17485.
40. **Lerner A**, Soto J and Rosen J. Chemotherapy as treatment for colocolonic intussusception associated with acquired immunodeficiency-related lymphoma. Submitted.
41. Dong, H, Carlton M, **Lerner A** and Epstein PM. Effect of cAMP signaling on expression of glucocorticoid receptor, Bim and Bad in glucocorticoid-sensitive and resistant multiple myeloma cells. In preparation.
42. Sinha A, Faller DV, **Lerner A** and Denis GV. Aggressive B1-like lymphomas of E $\mu$ -BRD2 transgenic mice lack immunoglobulin gene diversity. In preparation.

## Support

RO1 CA106705 (Lerner) NIH/NCI cAMP-mediated apoptosis in lymphoid malignancies The major goals of this project are to elucidate the mechanisms by which PDE4 inhibitors induce apoptosis in B-CLL cells and to contrast the effects of EPAC and PKA signaling in B-CLL.	07/01/05 – 06/30/10	40%
RO1 CA114094-01 (Lerner) NIH/NCI Analysis of AND-34 in human breast cancer The major goal of this project is to test the mechanism by which over-expression of AND-34 induces PI3K and Rac activation and anti-estrogen resistance in breast cancer cell lines.	07/01/05-06/30/09	20%
T32 HL007501-25 (Lerner) Research Training in Blood Diseases and Resources This NHLBI-supported Hematology Training Program supports four pre-doctoral and four post-doctoral students/year.	07/01/03-06/30/08	15%
No grant number (Lerner) EXHIBIT A	1/1/06 – 12/31/08 3	5%

Logica Foundation

Comparative studies of NSP family members

The major goal of this project is to contrast the role of three NSP family members (NSP-1, AND-34/BCAR3, and NSP-3/CHAT/SHEP1) in hematopoietic tissues, breast tumors and during development.